

Figure 1, HCV J4L6 genome wild-type cDNA sequence, reference accession number  
AF054247,

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9541 ccgtgagccg catgactgca gagagtgtg atactggcct ctctgcagat catgt

**Figure 2, codon optimised HCV Core polynucleotide**

ATGAGCACCAACCCCAAGCCCCAGCGCAAGACCAAGCGGAACACCAACCGGAGACCCAGGA  
CGTCAAGTTCCCAGGAGGAGGCCAGATCGTGGGCGGCGTGTACCTGCTGCCCCGCCGGGGGC  
CCCGGCTGGGCGTGCGCGCCACCCGCAAGACCAGCGAGCGCTCCCAGCCAAGAGGCAGACGC  
CAGCCGATCCCGAAGGCCCGCCGCCCTGAGGGCCGGGCTTGGGCCCAGCCAGGCTACCCCTG  
GCCCCGTGTATGGCAACGAGGGCCTGGGATGGGCTGGGTGGCTCCTCAGCCCCCGGGGGTCTA  
GGCCCAGTTGGGGACCGACCGACCCCGCAGGCGCAGCCGCAACCTGGGAAAGGTGATCGAC  
ACGCTCACCTGCGGCTTCGCCGACTTGATGGGATACATCCCTCTGGTGGGGGCCCTCTGGG  
CGGAGCCGCGCGCGCCCTGGCTCACGGGGTCCGGGTGCTCGAGGACGGGGTGAACCTACGCCA  
CCGGGAACCTGCCCCGGCTGCAGCTTCTCCATCTTCCTGCTGGCGCTGCTGAGCTGCCTCACC  
ATCCCCGCTAGCGCATGA

**Figure 3, Codon optimised HCV NS3 polynucleotide**

ATGGCCCCCATCACCGCCTACAGCCAGCAGACCCGGGGACTGCTCGGCTGCATCATCACCTC  
TCTGACAGGCCGGGATAAGAACCAGGTGGAGGGCGAGGTGCAGGTCGTCTCGACCGCTACCC  
AAAGCTTCCTGGCCACCTGTATCAACGGAGTCTGCTGGACGGTGTACCATGGCGCCGGCAGC  
AAGACCCTCGCCGGGCCTAAGGGCCCCATCACCCAGATGTACACCAACGTGGACCAGGACCT  
GGTGGGCTGGCAGGCGCCCCCGGGGCGAGGAGTATGACCCCATGCACCTGCGGGAGCTCTG  
ACCTGTATCTGGTGACCAGACATGCCGATGTCATCCCGGTGAGGCGTCGCGGGGACAGTAGA  
GGGAGCCTGCTGAGCCCCCGCCCCGTCAGCTACCTGAAGGGGTCCGTGGGCGGCCCCCTGCT  
GTGCCCCCTCTGGCCACGTGGTCGGCATCTTCAGGGCCGCGGTGTGCACGCGCGGCGTGGCCA  
AGGCCGTGGACTTTATCCCCGTGGAGAGCATGGAGACCACCATGCGCTCCCCCGTGTTTACC  
GACAACAGCAGCCCCCCCCGCGGTGCCTCAGACCTTCCAGGTCGCCCACCTCCATGCTCCGAC  
GGGCTCCGGGAAGTCCACGAAGGTGCCCGCCGCGTACGCGGCCAGGGATACAAGGTGCTGG  
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GCCAGGAGTGCCACTCTACCGACAGCACCACAATCCTGGGCATCGGCACCGTCCCTCGACCAG  
GCTGAGACAGCGGGCGCCCGCCTGGTGGTGCTGGCCACGGCCACTCCCCCGGCTCCGTAC  
GGTGCCCCACCCCAATATCGAGGAGGTGGCCCTGAGCAACAACGGCGAGATCCCATTCTACG  
GCAAGGCTATCCCGATCGAGGCGATTAAGGGAGGCAGACATCTGATCTTCTGCCACAGCAAG  
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CCGGCCTGACACACATCGACGCCCATTTCTTGTCCCAAACCAAGCAGGCTGGCGACAATTC  
CCGTATCTGGTCGCGTACCAGGCCACGGTGTGCGCGCGTGCAGGCTCCCCCCCCCTAGCTG  
GGATCAGATGTGGAAGTGCTGATCCGCCTGAAGCCACCTGCATGGGCCCCCCCCCTGC  
TGTACCGCCTGGGCGCGGTGCAGAACGAAGTCACCTTGACCCACCCCATCACCAAGTACATC

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ATGGCGTGTCATGTCCGCTGACCTGGAGGTGGTCACCTGA



**Figure 4, codon optimised HCV NS4B polynucleotide**

ATGTTTTGGGCCAAGCATATGTGGAAC TTCATCAGCGGCATCCAGTACCTCGCCGGGCTGAG  
CACCTCCCGGGCAACCCCGCGATCGCAAGCCTGATGGCGTTCACAGCGAGCATCACCTCCC  
CCCTGACTACCCAGAACACACTGCTGTTCAACATCCTGGGGGGCTGGGTGCGCCGCTCAGCTG  
GCCCCCTCCTTCCGCCGCCAGCGCCTTTGTGGGGGCGGGAATCGCCGGGGCCGCCGTCGGCTC  
CATCGGACTGGGCAAGGTGCTGGTCGACATCCTGGCGGGCTACGGCGCGGGAGTCGCCGGAG  
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GCGCCACGTGGGCCCCGGGCGAGGGAGCCGTGCAGTGGATGAACCGCCTGATCGCCTTTGCCT  
CCCGCGGCAACCACGTCAGCCCTACACATTACGTGCCCCGAGAGCGATGCCGCCGCCCGCGTG  
ACCCAGATCCTGAGCTCCCTGACCATCACCCAGCTGCTCAAGAGGCTGCACCAGTGGATCAA  
CGAGGACTGCTCCACCCCTTGCTGA

**Figure 5, codon optimised HCV NS5B polynucleotide**

ATGTCCATGTCCTACACCTGGACCGGCGCCCTGATCACCCCCTGCGCCGCCGAGGAGAGCAA  
GCTCCCGATTAACCCCCTGTCCAACCTCTCTGCTCCGCCATCACAAACATGGTGTATGCCACCA  
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GTGATGGGGAGTTCTACGGCTTCCAGTACAGCCCGAAGCAGAGGGTGGAGTTCTTGGTGAA  
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GGAGGACGCGCGGCCCTGCGGGCGTTCACAGAGGCCATGACACGCTACAGTGTCCCCCCCCG  
GCGACCCCCCCCAGCCCGAATACGATCTGGAGCTCATCACTAGTTGCAGCTCGAACGTGTCT  
GTGGCCCATGACGCTTCTGGCAAACGGGTGTATTATCTGACGCGCGATCCCACCACCCCCCT  
CGCCAGAGCCGCGTGGGAGACAGCTCGGCACACCCCTGTGAACTCTTGGCTGGGCAACATCA  
TCATGTACGCCCCTACCCTGTGGGCTCGCATGATCCTGATGACCCACTTCTTCAGTATCCTC  
CTCGCTCAGGAGCAGCTGGAGAAGGCGCTCGACTGCCAGATCTACGGCGCCTGCTATAGTAT  
CGAGCCTCTCGACCTGCCCCAGATCATCGAGAGACTGCATGGGCTCAGCGCCTTCTCCCTCC  
ATAGTTACTCTCCTGGAGAAATTAACCGGGTGGCGAGCTGTCTGCGGAAGCTCGGCGTCCCC  
CCTCTGCGCGTTTGGCGGCATCGCGCCAGGAGTGTGAGGGCCAAGCTGCTGAGCCAGGGCGG  
AAGGGCCGCCACCTGCGGCCGGTATCTCTTCAACTGGGCCGTGCGCACCAAGCTCAAGCTCA  
CCCCATCCCTGCCGCCAGTCAGCTGGATCTCAGTGGGTGGTTTCGTGGCCGGCTATTCTGGC  
GGCGACATCTACCACTCCCTCAGCAGGGCGCGCCCCCGCTGGTTCCCCCTGTGCCTGCTGCT  
CCTGAGCGTCGGAGTCGGCATCTACCTGCTGCCCAACCGCTGA

Figure 6, Translation of HCV J4L6 genome (wild-type sequence)

1 MSTNPKPQRK TKRNTNRRPQ DVKFPGGGQI VGGVYLLPRR GPRLGVRATR KASERSQPRG  
61 RRQPIPKARR PEGRAWAQPG YPWPLYGNEG LGWAGWLLSP RGSRPSWGPT DPRRRSRNLG  
121 KVIDTLTCGF ADLMGYIPLV GAPLGGAARA LAHGVRVLED GVNYATGNLP GCSFSIFLLA  
181 LLSCLTIPAS AYEVRNVSGI YHVTNDCSNS SIVYEAADVI MHTPGCVPCV QEGNSSRCWV  
241 ALTPTLAARN ASVPTTTIRR HVDLLVGTA A FCSAMYVGDL CGSIFLVSQ L FTFSPRRHET  
301 VQDCNCSIYP GHVSGHRMAW DMMNWSPTT ALVVSQLLRI PQAVVDMVAG AHWGVLAGLA  
361 YYSVMGNWAK VLIvallFAG VDGETHTTGR VAGHTTSGFT SLFSSGASQK IQLVNTNGSW  
421 HINRTALNCN DSLQTGFFAA LFYAHKFNSS GCPERMASCR PIDWFAQWG PITYTKPNSS  
481 DQRPYCWYHA PRPCGVVPAS QVCGPVYCF T PSPVVVGTTD RSGVPTYSWG ENETDVMLLN  
541 NTRPPQGNWF GCTWMNSTGF TKTCGGPPCN IGGVGNRTLI CPTDCFRKHP EATYTKCGSG  
601 PWLTPRCLVD YPYRLWHYPC TLNFSIFKVR MYVGGVEHRL NAACNWRGE RCNLED RDRS  
661 ELSPLLLSTT EWQILPCAFT TLPALSTGLI HLHQNI DVQ YLYGVGSAFV SFAIKWEYIL  
721 LLFLLLADAR VCACLWMLL IAQAEAALEN LVVLNAASVA GAHGILSFLV FFCAAWYIKG  
781 RLAPGAAYAF YGVWPLLLLL LALPPRAYAL DREMAASC GG AVLVLGLVFLT LSPYYKVFLT  
841 RLIWWLQYFI TRAEAHMQVW VPPLNVRGGR DAIILLTCAV HPELIFDITK LLLAILGPLM  
901 VLQAGITRVP YFVRAQGLIR ACMLVRKVAG GHYVQMVFMK LGALTGT YVY NHLTPLRDWA  
961 HAGLRDLAVA VEPVVFSA ME TKVITWGADT AACGDIILGL PVSARRGKEI FLGPADSLEG  
1021 QGWRL LAPIT AYSQQTRGVL GCIITSLTGR DKNQVEGEVQ VVSTATQ SFL ATCINGVCWT  
1081 VYHGAGSKTL AGPKGPITQM YTNVDL DLVG WQAPPGARSM TPCSCGSSDL YLVTRHADVI  
1141 PVRRRGDSRG SLLSPRPVS Y LKGSSGGPLL CPSGHVVG VF RAAVCTR GVA KAVDFIPVES  
1201 METTMRSPVF TDNSTPPAVP QTFQVAHLHA PTGSGKSTKV PAAYAAQGYK VLVLNPSVAA  
1261 TLGFGAYMSK AHGIDPNIRT GVRTITTGGS ITYSTYKFL ADGGC SGGAY DIIICDECHS  
1321 TDSTTILGIG TVLDQAETAG ARLVVLATAT PPGSVTV PHP NIEBIGLSNN GEIPFYGKAI  
1381 PIEAIKGRH LIFCHSKK KC DELAAKL TGL GLNAVAYYRG LDVSVIPP IGV DVVVVATDAL  
1441 MTGFTGDFDS VIDCNTCVTQ TVDFS LDPTF TIETTTVPQD AVSR SQRRGR TGRGRSGIYR  
1501 FVTPGERPSG MFDSSVLCEC YDAGCAWYEL TPAETSVRLR AYLNTPGLPV CQDHLEFWES  
1561 VFTGLTHIDA HFLSQTKQAG DNFPYLVAYQ ATVCARAQAP PPSWDQM WKC LIRLKP TLHG  
1621 PTPLLYRLGA VQNEVILTHP ITKYIMACMS ADLEVVTSTW VLVGGVLAAL AAYCLTTGSV  
1681 VIVGRIILSG KPAVVPDREV LYQEFDEMEE CASQLPYIEQ GMQLAEQFKQ KALGLLQTAT  
1741 KQAEAAAPVV ESKWRALETF WAKHMWNFIS GIQYLAGLST LPGNPAIASL MAFTASITSP  
1801 LTTQNTLLFN ILGGWVAAQL APPSAASAFV GAGIAGAAVG SIGLGKVLVD ILAGYGAGVA  
1861 GALVAFK VMS GEVPSTEDLV NLLPAILSPG ALVVG VVCAA ILRRHVGPGE GAVQWMNRLI  
1921 AFASRGNHVS PTHYVPESDA AARVTQILSS LTITQLLKRL HQWINE DCST PCSG SWLRDV  
1981 WDWICTVLTD FKTWLQSKLL PRLPGVPFLS CQRGYKGVWR GDGIMQTTCP CGAQIAGHV K  
2041 NGSMRIVGPR TCSNTWHGTF PINAYTTGPC TPSPAPNYSR ALWRVAAEEY VEVTRVGDFH  
2101 YVTGMTTDNV KCPCQVPAP E FFTEVDGVRL HRYAPACKPL LREDVTFQVG LNQYL VGSQ L

2161 PCEPEPDVTV LTSMLTDPSH ITAETAKRRL ARGSPPSLAS SSASQLSAPS LKATCTTHHD  
2221 SPDADLIEAN LLWRQEMGGN ITRVESENKV VILDSFEPLH AEGDEREISV AAEILRKSRL  
2281 FPSALPIWAR PDYNPPLLES WKDPDYVPPV VHGCPLPPTK APPIPPPRRK RTVVLTESNV  
2341 SSALAEELATK TFGSSGSSAV DSGTATALPD LASDDGDKGS DVESYSSMPP LEGEPGDPDL  
2401 SDGSWSTVSE EASEDVVCCS MSYTWGTALI TPCAAEESKL PINPLSNSLL RHHNMVYATT  
2461 SRSASLRQKK VTFDRLQVLD DHYRDVLKEM KAKASTVKAK LLSIEEACKL TPPHSAKSKF  
2521 GYGAKDVRNL SSRVNHIRS VWEDLLEDTE TPIDTTIMAK SEVFCVQPEK GGRKPARLIV  
2581 FPDLGVRVCE KMALYDVVST LPQAVMGSSY GFQYSPKQRV EFLVNTWKSK KCPMGFSYDT  
2641 RCFDSTVTES DIRVEESIQ CCDLAPEARQ AIRSLTERLY IGGPLTNSKG QNCGYRRCRA  
2701 SGVLTTS CGN TLTCYLKATA ACRAAKLQDC TMLVNGDDL VICESAGTQE DAAALRAFTE  
2761 AMTRYSAAPP DPPEYDLE LITSCSSNVS VAHDASGKRV YYLTRDPTTP LARAAWETAR  
2821 HTPINSWLGN IIMYAPTLWA RMILMTHFFS ILLAQEQLEK ALDCQIYGAC YSIEPLDLPO  
2881 IIERLHGLSA FTLHSYSPGE INRVASCLRK LGVPPLRTWR HRARSVRACL LSQGGRAATC  
2941 GRYLFNWAVR TKLKLTPIPA ASQLDLGWF VAGYSGGDIY HSLSRARPRW FPLCLLLLSV  
3001 GVGIIYLLPNR

Figure 7, p7313-ie

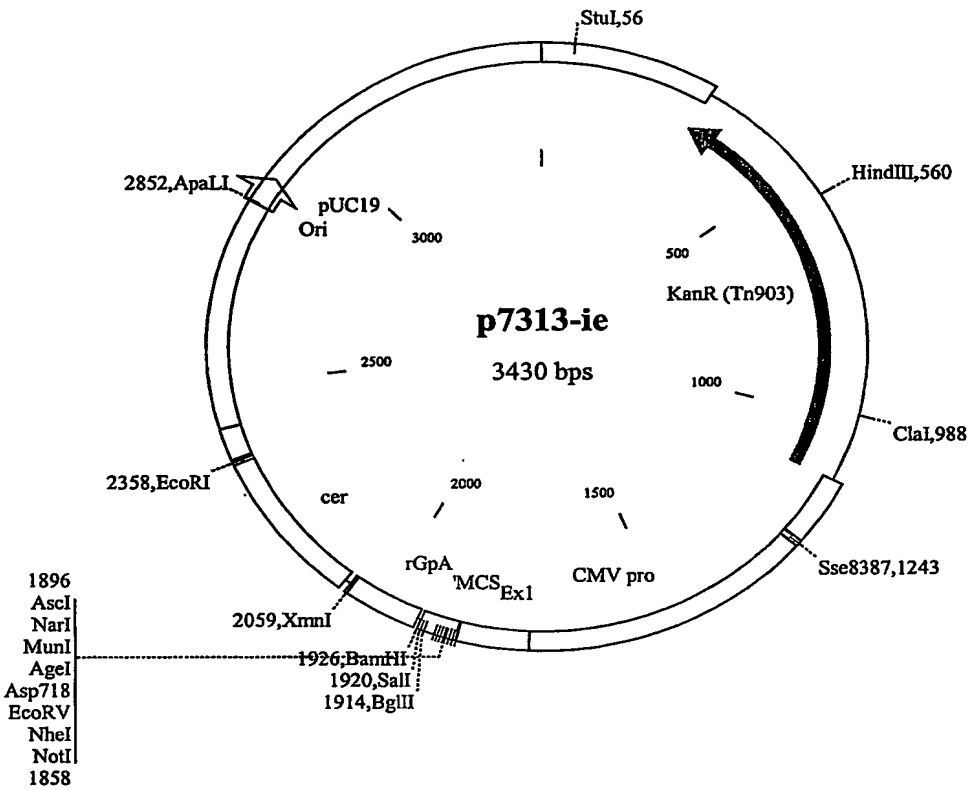


Figure 8, Immune responses to Core

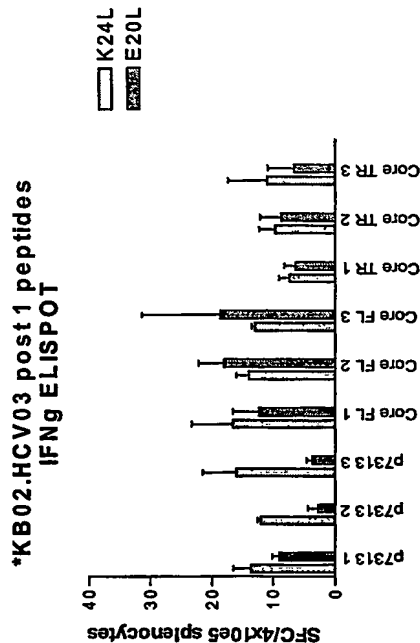
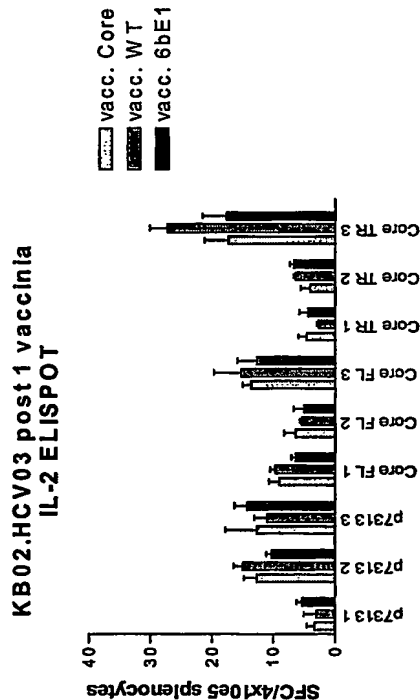
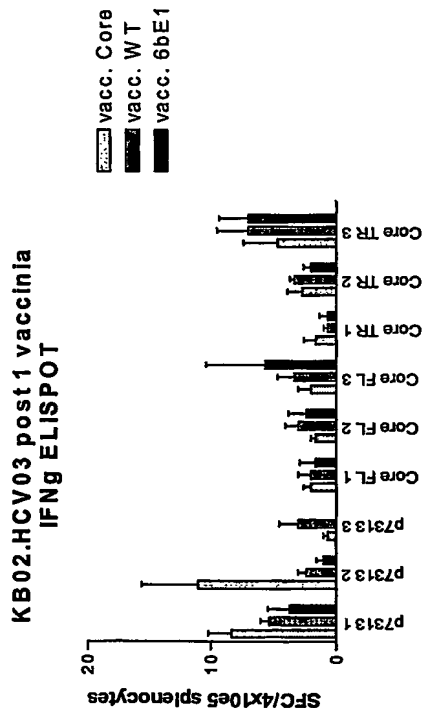
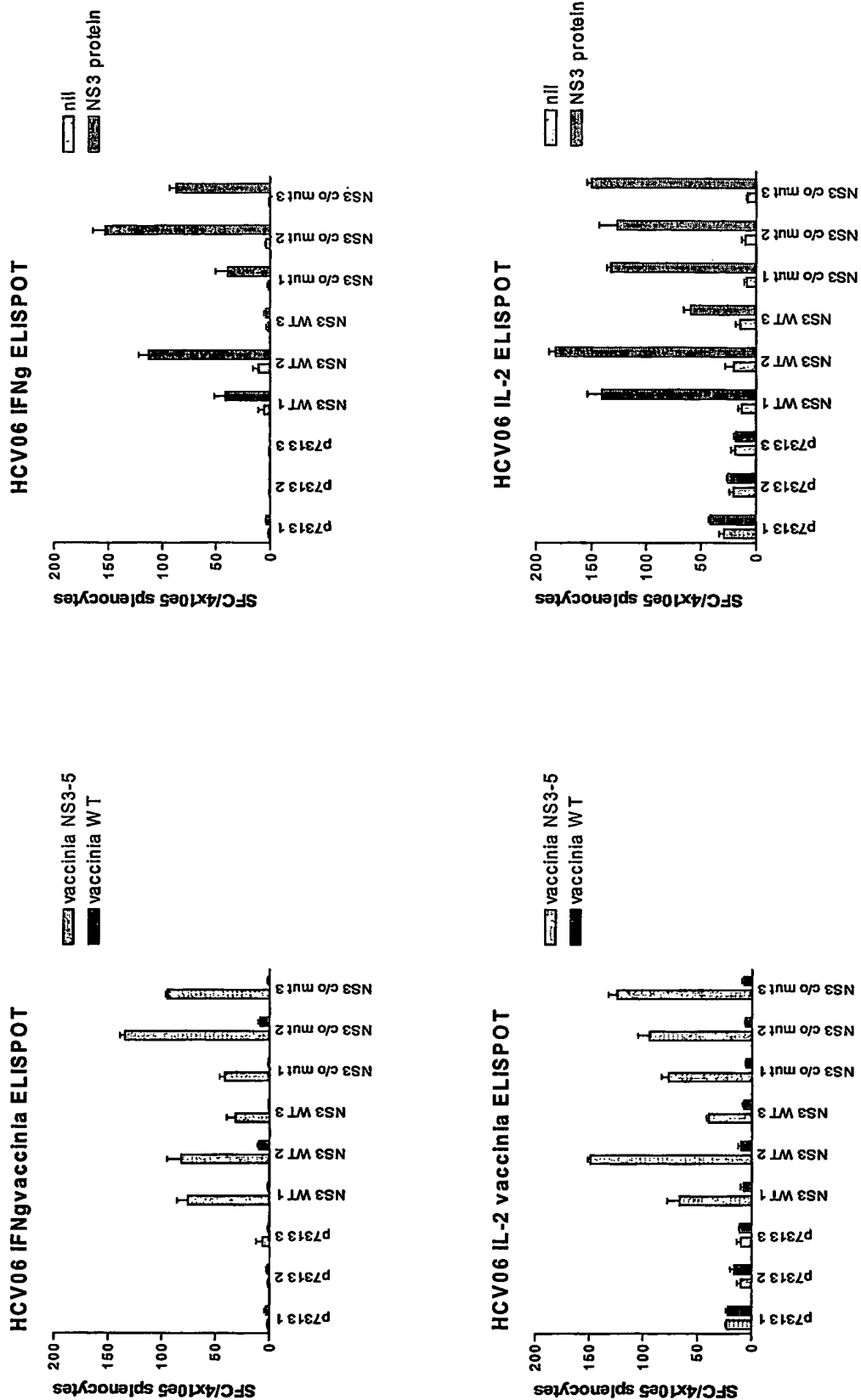


Figure 9, NS3 immunogenicity



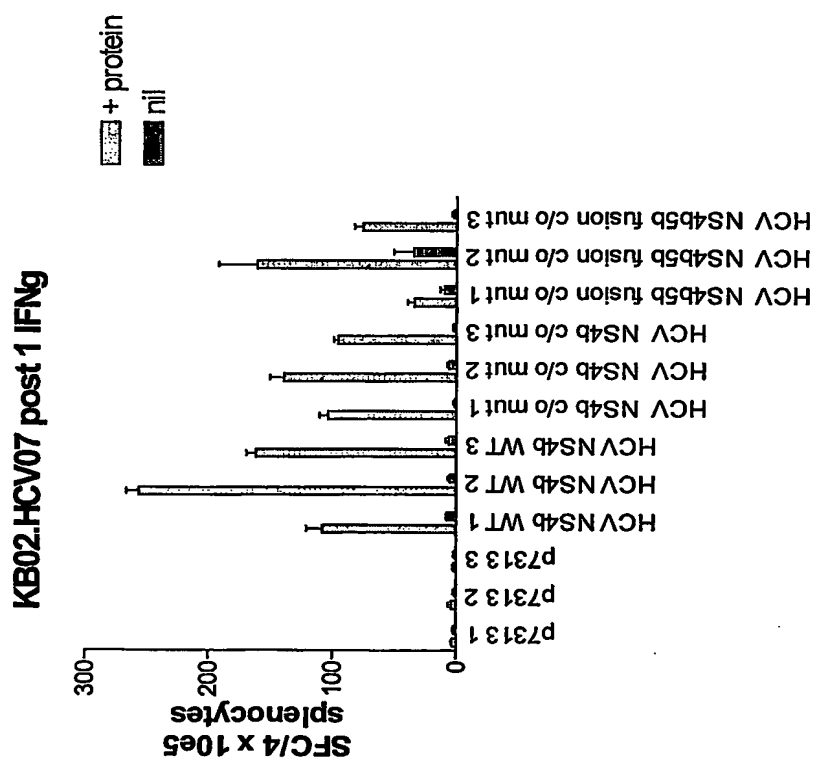
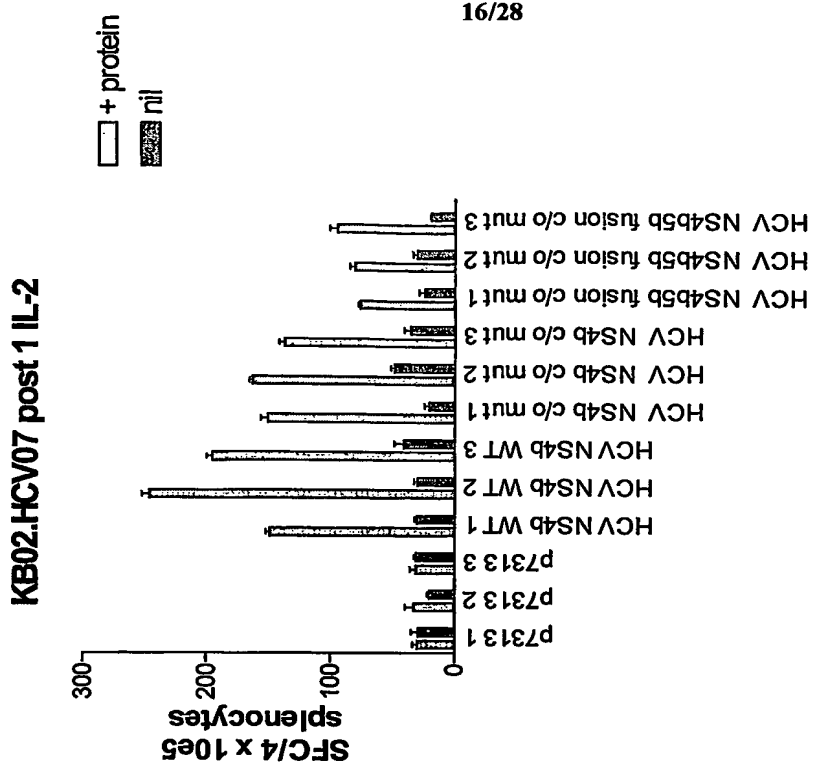


Figure 10, Immune responses to NS4B



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Figure 11, NS5B immune responses

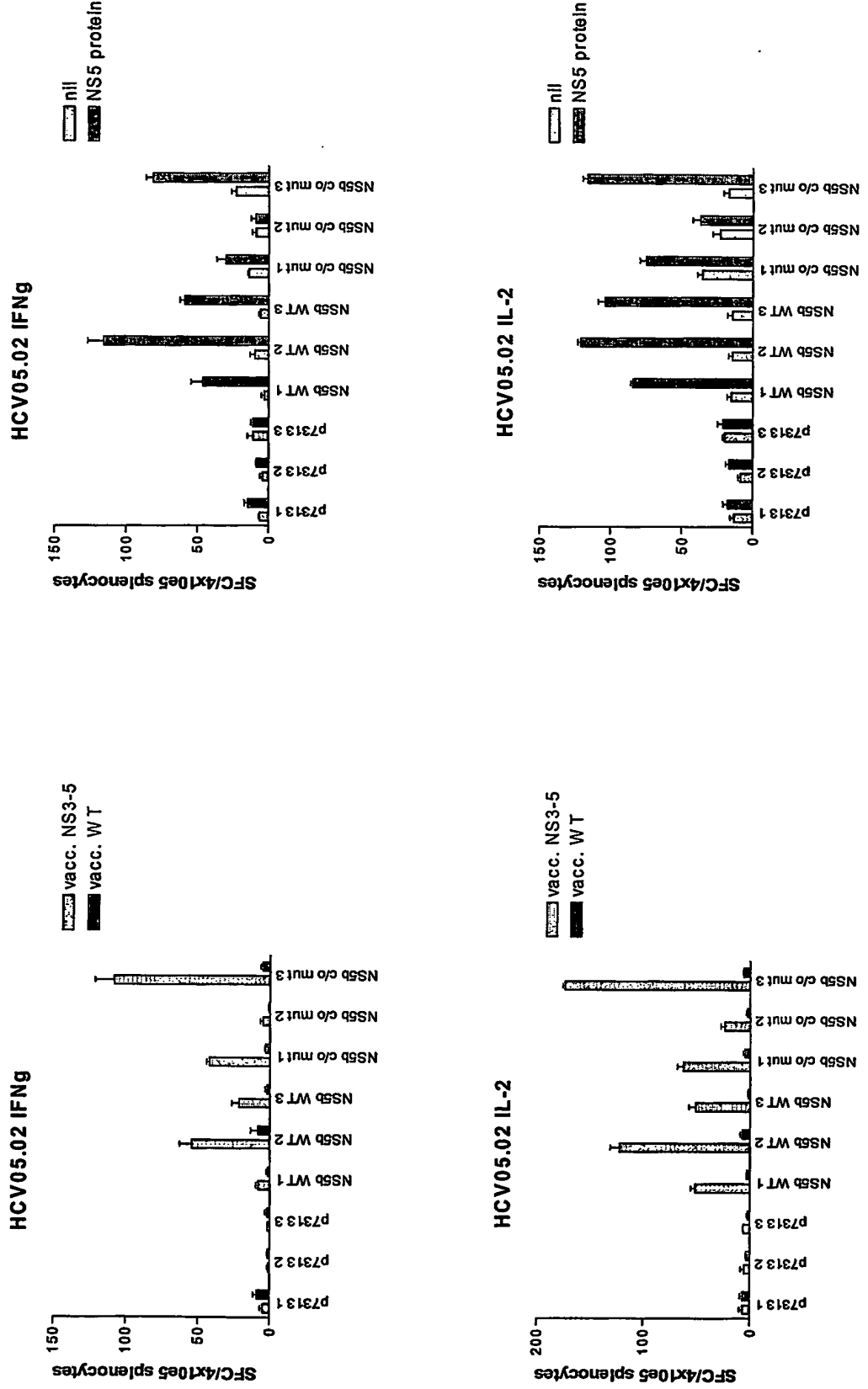


FIG. 12

# Anti-HCV NS5B

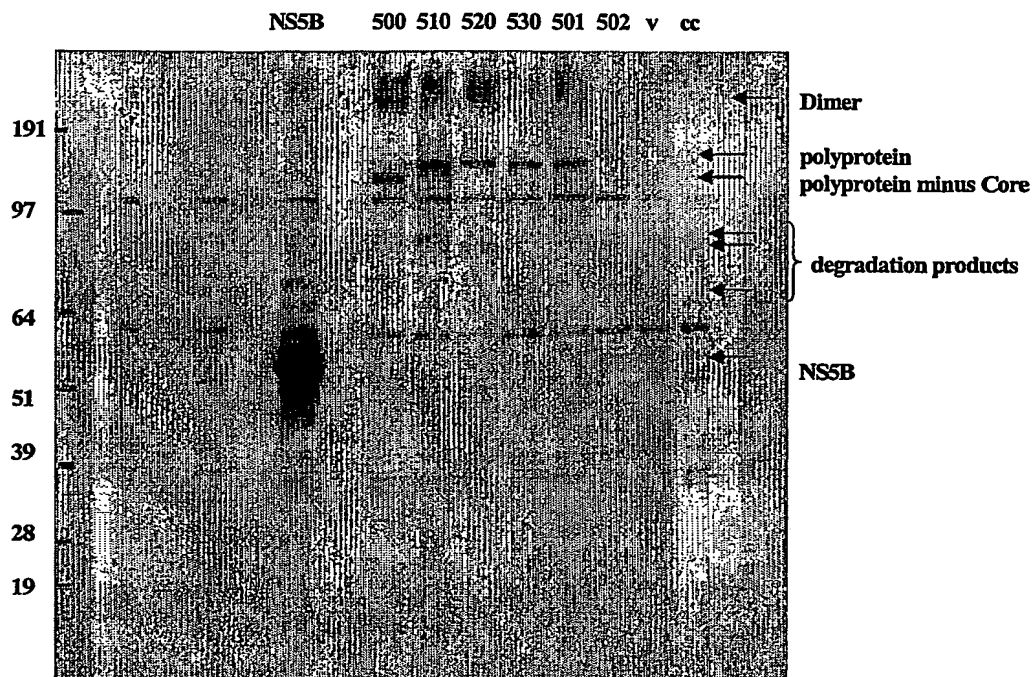
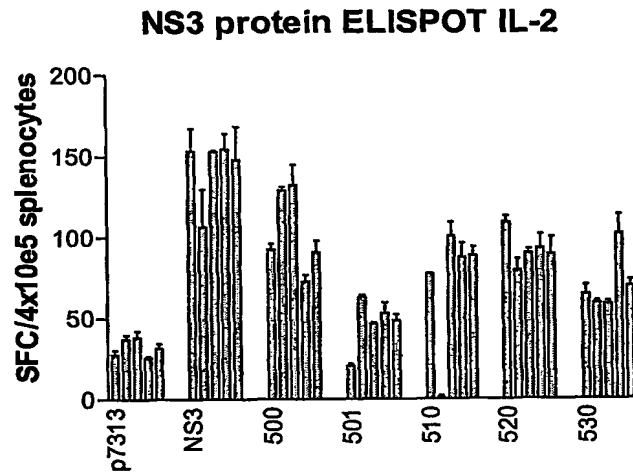
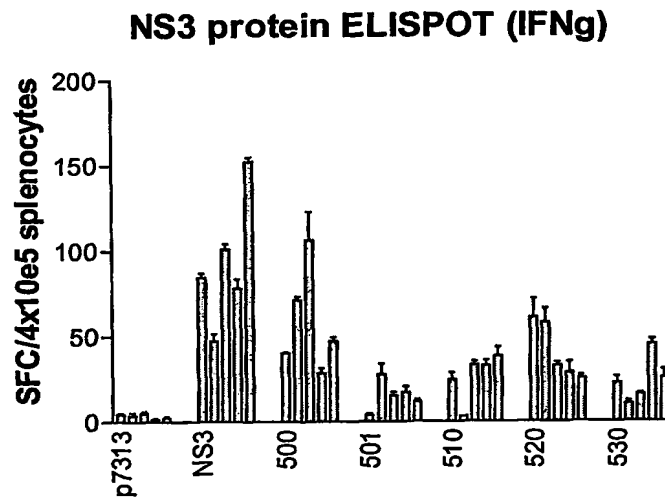


FIG. 13, A



B.



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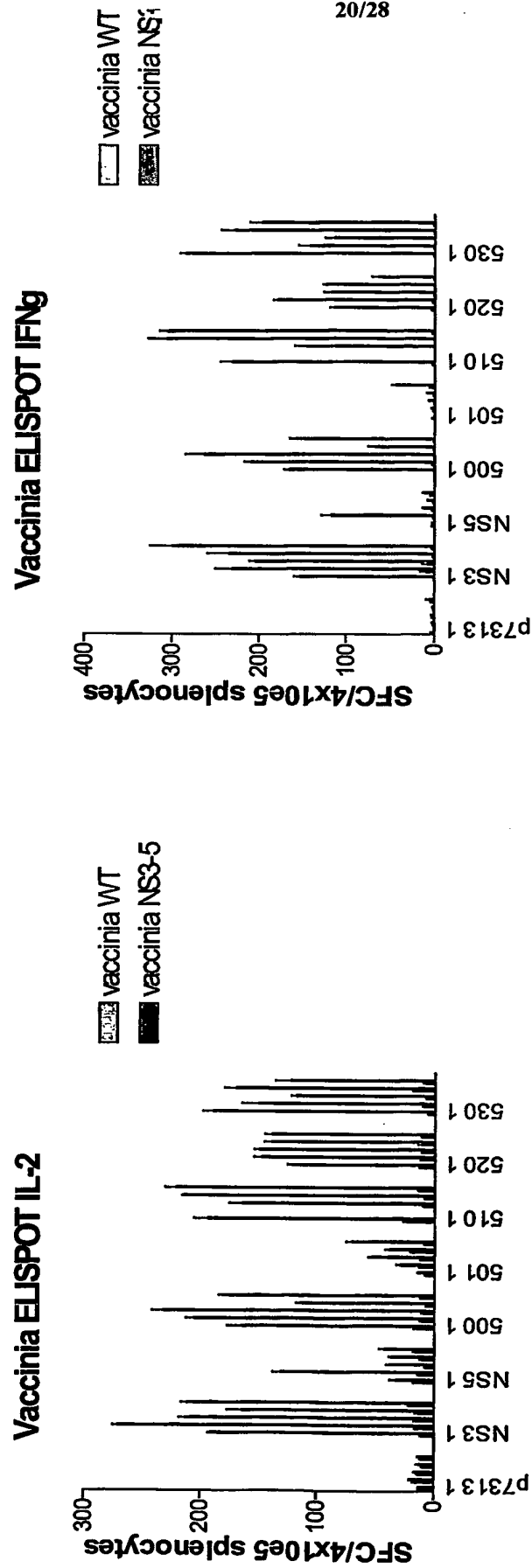


FIG 14.

FIG. 15,

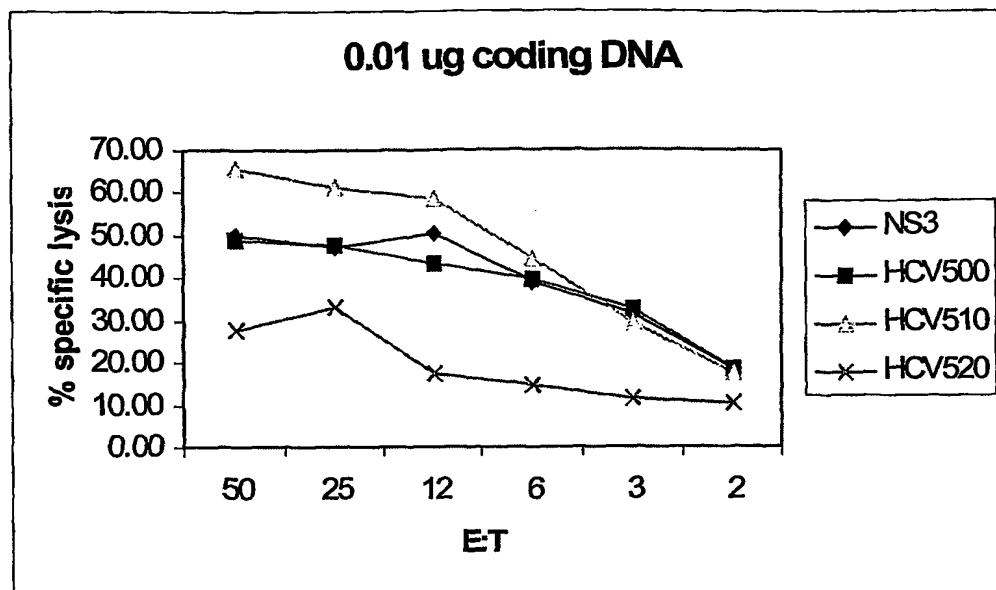


FIG. 16,

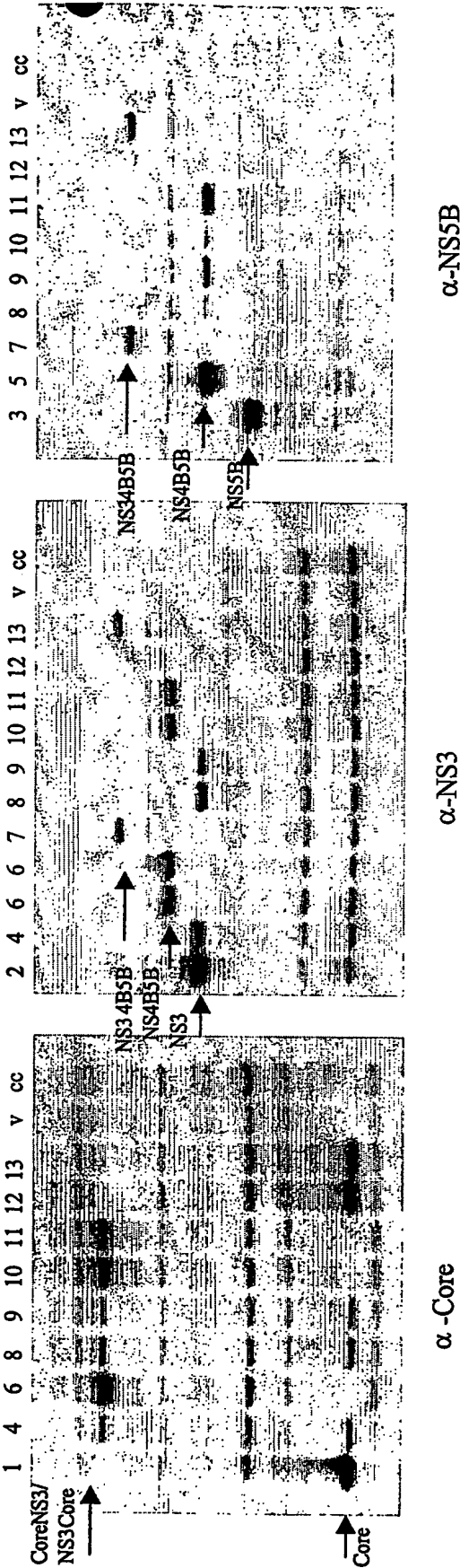


FIG. 17, Comparison of NS3 T cell response induced by dual promoter constructs.

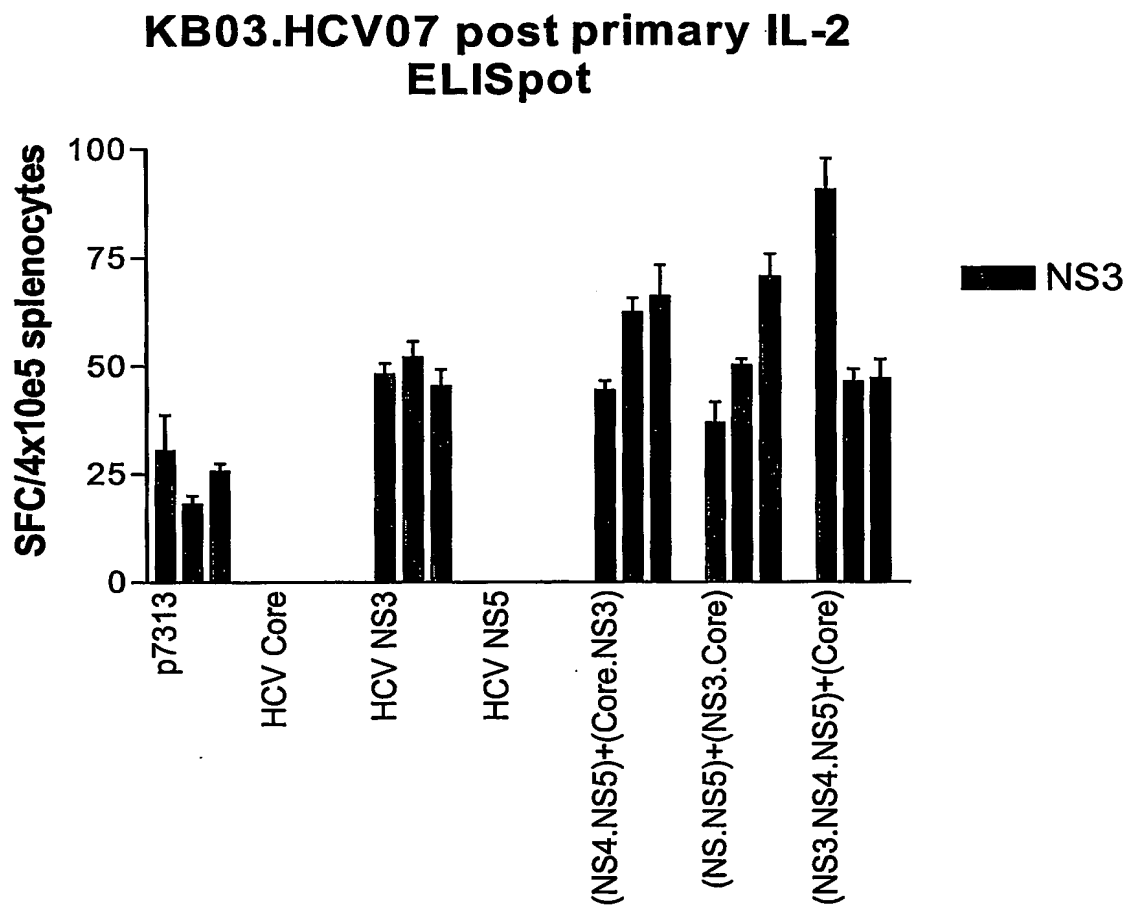
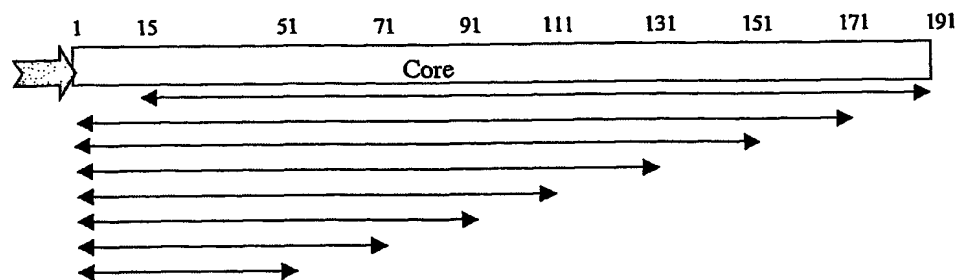


FIG. 18,



MW C191 CA15 C171 C151 C131 C111 C91 C71 C51



FIG. 19.

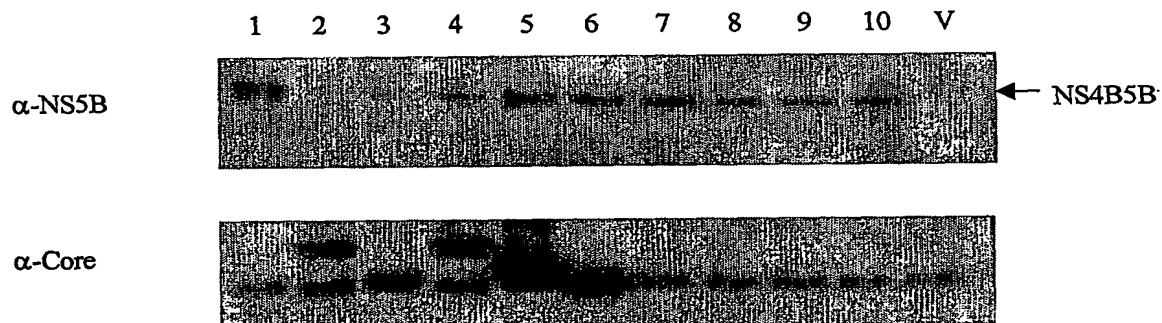
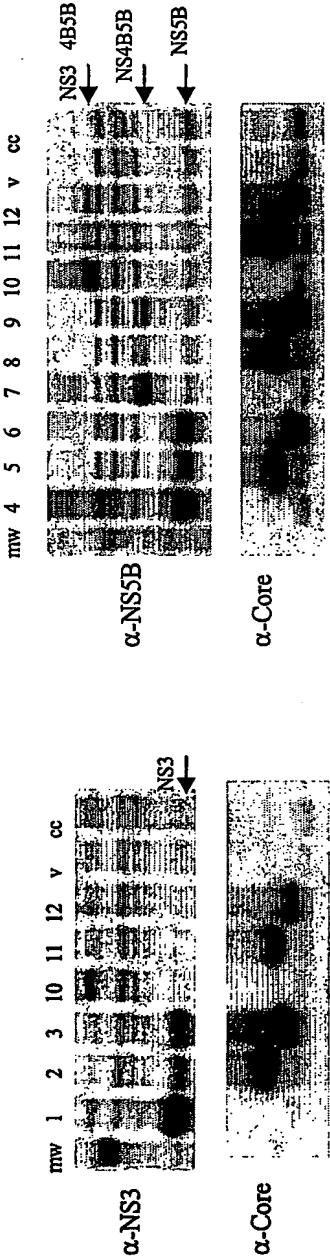




FIG. 20,

Effect of Core and Core<sub>151</sub> upon expression of NS3, NS5B, NS4B5B, and NS34B5B after co-transfection in 293T cells



Samples:

1. p7/NS3 + v
2. p7/NS3 + p7/Core
3. p7/NS3 + p7/Core151
4. p7/NS5B + v
5. p7/NS5B + p7/Core
6. p7/NS5B + p7/Core151
7. p7/NS4B5B + v
8. p7/NS4B5B + p7/Core
9. p7/NS4B5B + p7/Core151
10. p7/NS34B5B + v
11. p7/NS34B5B + p7/Core
12. p7/NS34B5B + p7/Core151

FIG. 21,

Effect on expression of fusion proteins, after substitution of Core<sub>1</sub> for Core<sub>191</sub>, in transient transfection in 293T cells

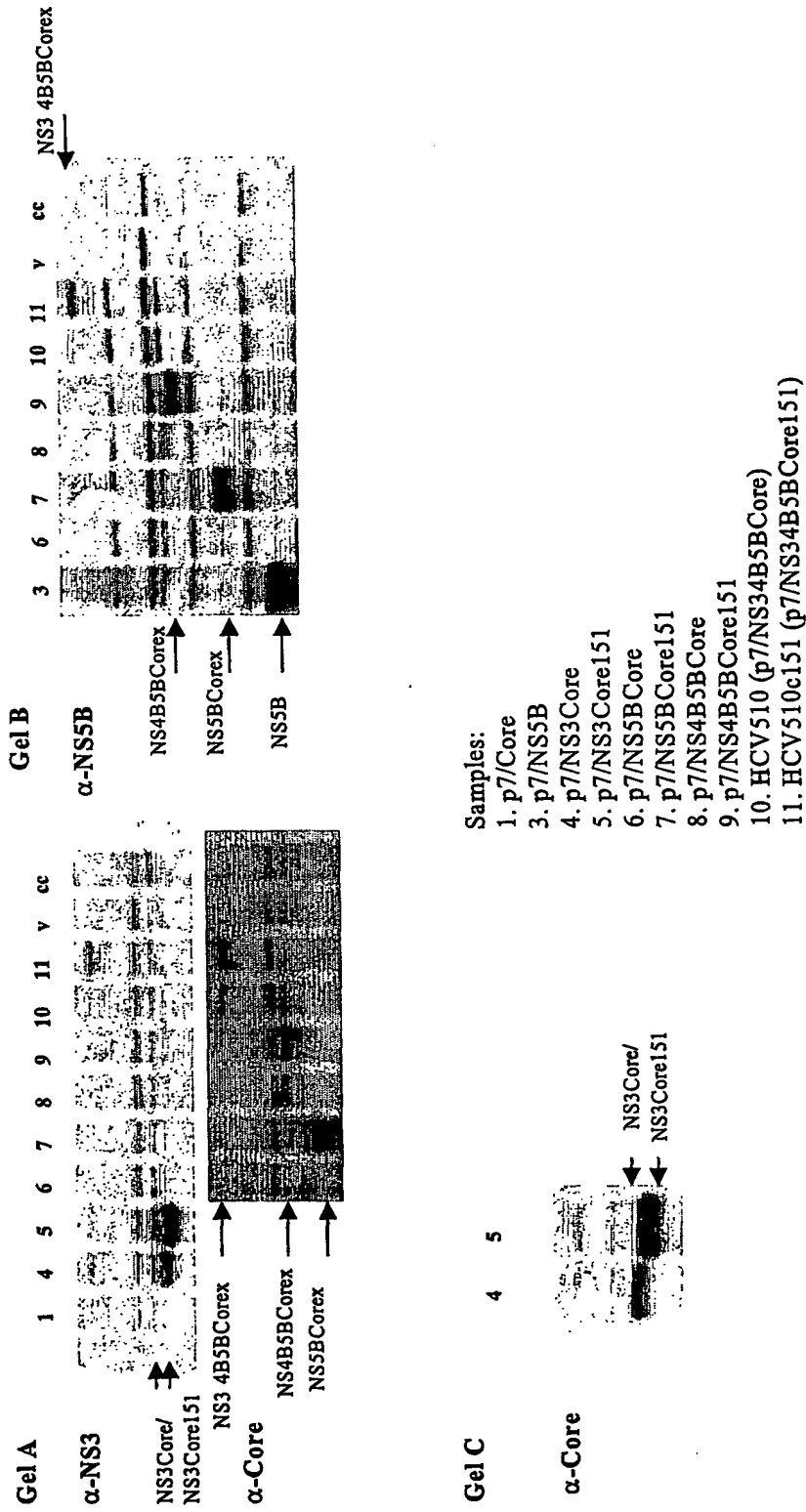
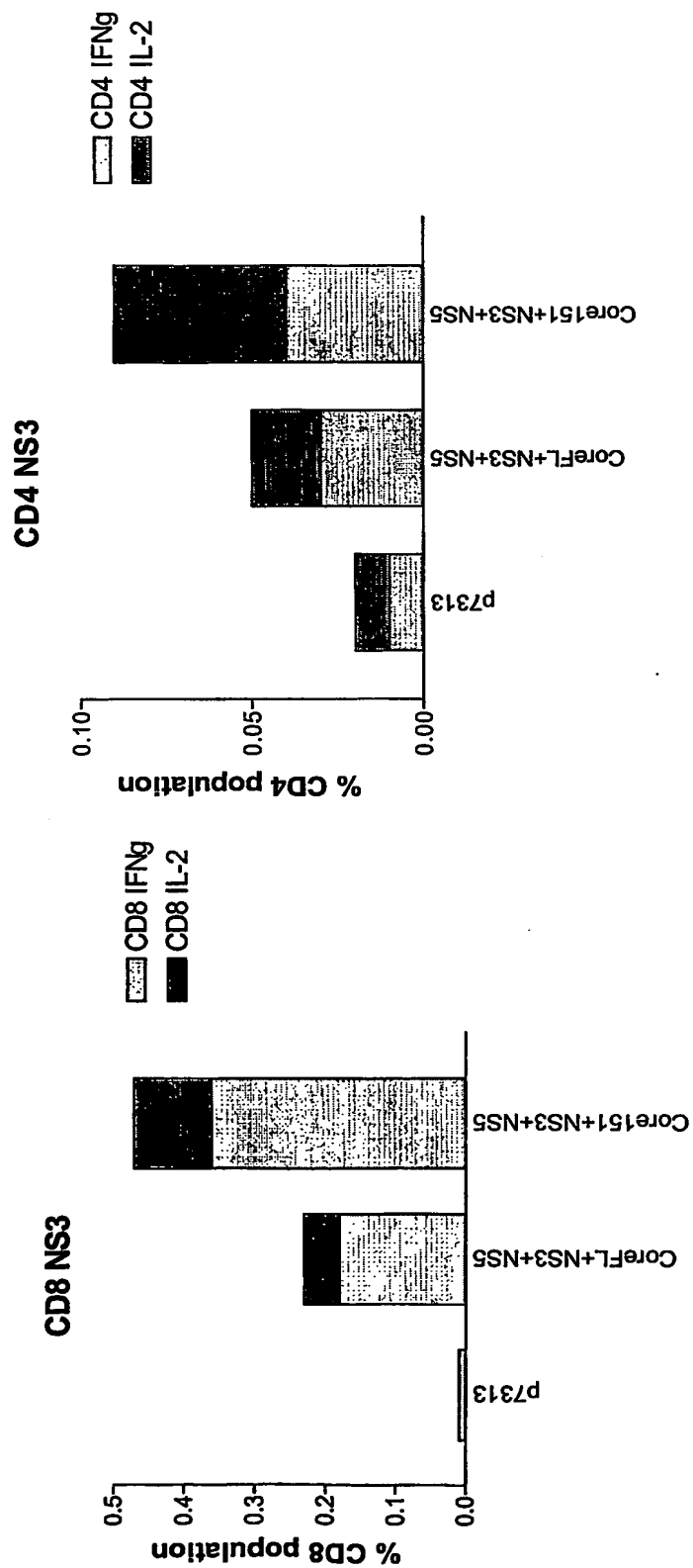


FIG. 22,



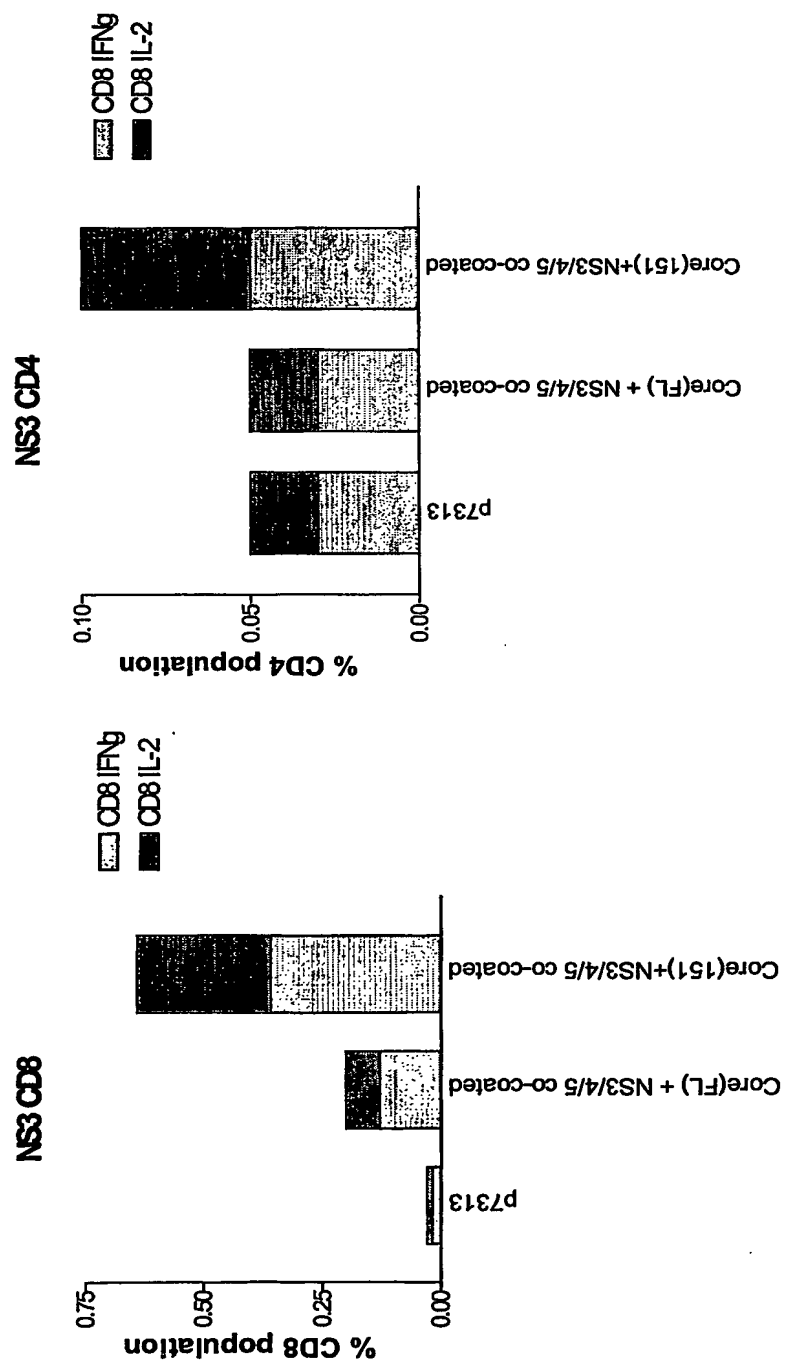


FIG. 23,

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